**PATENT** Short and Keller Attorney Docket No.: DIVER1280-3

Application No.: 09/685.432 Filed: October 10, 2000

Page 2

## Amendments to the Claims:

This listing of claims will replace all prior versions, and listings, of claims in the application.

Upon entry of the present amendment, the claims will stand as follows:

Please cancel claims 2, 13, 15, 18 and 26 without prejudice.

Please amend claims 1, 10, 16 and 19 as follows:

- 1. (Currently Amended) A method for identifying a bioactivity or a biomolecule an enzyme of interest, comprising:
  - obtaining a plurality of polynucleotides derived from a mixed population of (a) organisms or more than one organism;
  - normalizing the plurality of polynucleotides normalizing the representation of (b) organisms present in the plurality of polynucleotides to increase representation of rare species;
  - contacting a library containing clones of normalized polynucleotides from (b) (c) with at least one oligonucleotide probe labeled with a detectable molecule, wherein the probe comprises at least a portion of a polynucleotide sequence encoding an enzyme of interest;
  - incubating the clones under such conditions and for such time as to allow hybridization of complementary sequences; [[and
  - (d]]e) separating clones with an analyzer that detects the detectable molecule;
  - (f) contacting the separated clones with a reporter system that comprises a substrate for the enzyme of interest; and
  - identifying clones capable of modulating expression or activity of the reporter system thereby identifying a polynucleotide that encodes the enzyme of interest.

Claim 2. (Cancelled)

In re Application of:

Short and Keller

Application No.: 09/685,432 Filed: October 10, 2000

Page 3

Attorney Docket No.

Attorney Docket No.: DIVER1280-3

**PATENT** 

- 3. (Previously presented) The method of claim 1, wherein the library is an expression library.
- 4. (Previously presented) The method of claim 1, wherein the detectable molecule is a fluorescent molecule.
- 5. (Previously presented) The method of claim 1, wherein the analyzer is a FACS analyzer.
- 6. (Previously presented) The method of claim 1, wherein the mixed population of organisms is from an environmental sample.
- 7. (Previously presented) The method of claim 1, wherein the mixed population of organisms comprises microorganisms.
- 8. (Previously presented) The method of claim 6, wherein the environmental sample contains extremophiles.
- 9. (Previously presented) The method of claim 8, wherein the extremophiles are selected from the group consisting of hyperthermophiles, psychrophiles, halophiles, psychrotrophs, alkalophiles, and acidophiles.
- 10. (Currently Amended) The method of claim [[2]] 1, wherein the reporter system is a bioactive substrate.
- 11. (Previously presented) The method of claim 10, wherein the bioactive substrate comprises C12FDG.
- 12. (Previously presented) The method of claim 11, wherein the bioactive substrate further comprises a lipophilic tail.

In re Application of:

Short and Keller

Application No.: 09/685,432

Filed: October 10, 2000

Page 4

Claims 13-15. (Cancelled)

16. (Currently amended) The method of claim 15, wherein the <u>clones are encapsulated in a</u> microenvironment [[is]] selected from beads, high temperature agaroses, gel microdroplets, cells, ghost cells, macrophages, or liposomes.

PATENT

Attorney Docket No.: DIVER1280-3

17. (Previously presented) The method of claim 16, wherein the clones are encapsulated in a gel microdroplet.

Claim 18. (Cancelled)

19. (Currently Amended) The method of claim [[18]]1, wherein the enzyme is selected from the group consisting of lipases, esterases, proteases, glycosidases, glycosyl transferases, phosphatases, kinases, mono- and dioxygenases, haloperoxidases, lignin peroxidases, diarylpropane peroxidases, eposize hydrolases, nitrile hydratases, nitrilases, transaminases, amidases, and acylases.

20. (Previously presented) The method of claim 1, wherein the reporter system comprises a detectable label.

Claim 21. (Cancelled)

- 22. (Previously presented) The method of claim 1, wherein the polynucleotide of interest encodes a small molecule.
- 23. (Previously presented) The method of claim 1, wherein the polynucleotide of interest, or fragments thereof, comprise one or more operons, or portions thereof.

In re Application of:

Short and Keller Application No.: 09/685,432

Filed: October 10, 2000

Page 5

24. (Previously presented) The method of claim 23, wherein the operons, or portions thereof, encodes a complete or partial metabolic pathway.

**PATENT** 

Attorney Docket No.: DIVER1280-3

25. (Previously presented) The method of claim 24, wherein the operons or portions thereof encoding a complete or partial metabolic pathway encodes polyketide syntheses.

Claims 26-61. (Cancelled)